

SEQUENCE LISTING

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(1) GENERAL INFORMATION:

- (i) APPLICANT: CHARETTE, MARC F. RUEGER, DAVID C. HIGGINS, DENNIS
- (ii) TITLE OF INVENTION: ENHANCEMENT OF MORPHOGEN ACTIVITY
- (iii) NUMBER OF SEQUENCES: 9
- (iv) CORRESPONDENCE ADDRESS:
 (A) ADDRESSEE: PATENT ADMINISTRATOR, TESTA, HURWITZ & THIBEAULT, LLP STREET: 125 HIGH STREET

 - (C) CITY: BOSTON (D) STATE: MA (E) COUNTRY: USA (F) ZIP: 02110
- (V) COMPUTER READABLE FORM:

 (A) MEDIUM TYPE: Floppy disk

 (B) COMPUTER: IBM PC compatible

 (C) OPERATING SYSTEM: PC-DOS/MS-DOS
 - (D) SOFTWARE: PatentIn Release #1.0, Version #1.30
- (vi) CURRENT APPLICATION DATA:
 - (A) APPLICATION NUMBER:
 (B) FILING DATE:
 (C) CLASSIFICATION:
- (viii) ATTORNEY/AGENT INFORMATION:

 - (A) NAME: MEYERS, THOMAS C. (B) REGISTRATION NUMBER: 36,989
 - (C) REFERENCE/DOCKET NUMBER: CRP-158PR
 - (ix) TELECOMMUNICATION INFORMATION:
 - (A) TELEPHONE: (617) 248-7000 (B) TELEFAX: (617) 248-7100
- (2) INFORMATION FOR SEQ ID NO:1:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1822 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single

 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: cDNA
 - (vi) ORIGINAL SOURCE:

 - (A) ORGANISM: HOMO SAPIENS (F) TISSUE TYPE: HIPPOCAMPUS
 - (ix) FEATURE:
 - (1x) FEATURE:

 (A) NAME/KEY: CDS

 (B) LOCATION: 49..1341

 (C) IDENTIFICATION METHOD: experimental

 (D) OTHER INFORMATION: /function= "OSTEOGENIC PROTEIN"

 /product= "OP1"

 /evidence= EXPERIMENTAL
 /standard_name= "OP1"

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:
- GGTGCGGGCC CGGAGCCCGG AGCCCGGGTA GCGCGTAGAG CCGGCGCG ATG CAC GTG Met His Val
- CGC TCA CTG CGA GCT GCG GCG CCG CAC AGC TTC GTG GCG CTC TGG GCA Arg Ser Leu Arg Ala Ala Pro His Ser Phe Val Ala Leu Trp Ala 5 105
- CCC CTG TTC CTG CTG CGC TCC GCC CTG GCC GAC TTC AGC CTG GAC AAC Pro Leu Phe Leu Leu Arg Ser Ala Leu Ala Asp Phe Ser Leu Asp Asn 20 35 153
- GAG GTG CAC TCG AGC TTC ATC CAC CGG CGC CTC CGC AGC CAG GAG CGG 201



Glu	Val	His	Ser	Ser 40	Phe	Ile :	His	Arg	Arg 45	Leu .	Arg	Ser	Gln	Glu 50	Arg	
CGG Arg	GAG Glu	ATG Met	CAG Gln 55	CGC Arg	GAG Glu	ATC Ile	CTC Leu	TCC Ser 60	ATT Ile	TTG Leu	GGC Gly	TTG Leu	CCC Pro 65	CAC His	CGC Arg	249
				CTC Leu												297
				AAC Asn												345
				TCC Ser												393
				AGC Ser 120												441
				TTC Phe												489
			Tyr	CAC His												537
CCA Pro	GAA Glu 165	GGG Gly	GAA Glu	GCT Ala	GTC Val	ACG Thr 170	GCA Ala	GCC Ala	GAA Glu	TTC Phe	CGG Arg 175	ATC Ile	TAC Tyr	AAG Lys	GAC Asp	585
	Ile			CGC												633
CAG Gln	GTG Val	CTC Leu	CAG Gln	GAG Glu 200	CAC His	TTG Leu	GJ A GGC	AGG Arg	GAA Glu 205	TCG Ser	GAT Asp	CTC Leu	TTC Phe	CTG Leu 210	Leu	681
				Leu					Glu					Phe	GAC Asp	729
			Thr					Val					, His		CTG Leu	777
GGC Gly	CTG Leu 245	Glr	CTC Lev	TCG Ser	GTG Val	GAG Glu 250	Thr	CTC Leu	GAT Asp	GGG Gly	Glr 255	Ser	ATC Ile	AAC Asn	CCC Pro	825
	Let					Gly					Glr				CCC Pro 275	873
					Phe					. Val					ATC lle	921
				y Sei					r Glr					Thi	G CCC	969
			n Gl					Al					u Ası		C AGC r Ser	1017
		o Gl					Ly:					u Ty			c TTC r Phe	1065
	g As					n Asp					a Pr				C GCC r Ala 355	1113
															C ATG	1161



				360					365					3/0			
			AAC Asn 375													12	09
			GTG Val												GCC Ala	12	57
															AAA Lys	13	05
			ATG Met									TAG	CTCC	TCC		13	51
GAG	ATT	CAG .	ACCC!	rttg(GG G	CCAA	STTT'	r TC	TGGA'	TCCT	CCA'	TTGC'	TCG	CCTT	GGCCA	.G 14	11
GAA	CAG	CAG .	ACCA	ACTG	CC T	TTTG'	TGAG	A CC	TTCC	CCTC	CCT	ATCC	CCA	ACTT	TAAAG	G 14	71
TGT	GAGA	στα	TTAG	GAAA	CA T	GAGC.	AGCA'	r at	GGCT'	TTTG	ATC.	AGTT'	TTT	CAGT	GGCAG	c 15	31
ATC	CAAT	GAA	CAAG	ATCC	TA C	AAGC'	TGTG	C AG	GCAA	AACC	TAG	CAGG.	AAA	AAAA	AACAA	.C 15	91
GCA'	AAAT	GAA	TAAA	GGCC	GG G	CCAG	GTCA	T TG	GCTG	GGAA	GTC	TCAG	CCA	TGCA	CGGAC	т 16	551
CGT'	TTCC	AGA	GGTA	ATTA	TG A	.GCGC	CTAC	C AG	CCAG	GCCA	ccc	AGCC	GTG	GGAG	GAAGG	G 17	711
GGC	GTGG	CAA	GGGG	TGGG	CA C	ATTG	GTGT	C TG	TGCG	AAAG	GAA	TTAA	GAC	CCGG	AAGTT	c 17	771
CTG	TAAT.	AAA	TGTC	ACAA	TA A	AACG	AATG	A AT	GAAA	AAAA	AAA	AAAA	AAA	А		18	322

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(2) INFORMATION FOR SEQ ID NO:2:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 431 amino acids
 (B) TYPE: amino acid
 (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

Met His Val Arg Ser Leu Arg Ala Ala Ala Pro His Ser Phe Val Ala 1 1 15Leu Trp Ala Pro Leu Phe Leu Leu Arg Ser Ala Leu Ala Asp Phe Ser Leu Asp Asn Glu Val His Ser Ser Phe Ile His Arg Arg Leu Arg Ser 40 45Gln Glu Arg Arg Glu Met Gln Arg Glu Ile Leu Ser Ile Leu Gly Leu 50 60 Pro His Arg Pro Arg Pro His Leu Gln Gly Lys His Asn Ser Ala Pro 65 75 80 Met Phe Met Leu Asp Leu Tyr Asn Ala Met Ala Val Glu Glu Gly Gly 85 90 Gly Pro Gly Gly Gln Gly Phe Ser Tyr Pro Tyr Lys Ala Val Phe Ser 100 105 110 Thr Gln Gly Pro Pro Leu Ala Ser Leu Gln Asp Ser His Phe Leu Thr Asp Ala Asp Met Val Met Ser Phe Val Asn Leu Val Glu His Asp Lys 130 140 Glu Phe Phe His Pro Arg Tyr His His Arg Glu Phe Arg Phe Asp Leu 145 150 155 160Ser Lys Ile Pro Glu Gly Glu Ala Val Thr Ala Ala Glu Phe Arg Ile 165 170Tyr Lys Asp Tyr Ile Arg Glu Arg Phe Asp Asn Glu Thr Phe Arg Ile 180 180



Ser Val Tyr Gln Val Leu Gln Glu His Leu Gly Arg Glu Ser Asp Leu Phe Leu Leu Asp Ser Arg Thr Leu Trp Ala Ser Glu Glu Gly Trp Leu Val Phe Asp Ile Thr Ala Thr Ser Asn His Trp Val Val Asn Pro Arg 225 230 235 His Asn Leu Gly Leu Gln Leu Ser Val Glu Thr Leu Asp Gly Gln Ser 245 250 255 Ile Asn Pro Lys Leu Ala Gly Leu Ile Gly Arg His Gly Pro Gln Asn 260 270 Lys Gln Pro Phe Met Val Ala Phe Phe Lys Ala Thr Glu Val His Phe 275 280 285 Arg Ser Ile Arg Ser Thr Gly Ser Lys Gln Arg Ser Gln Asn Arg Ser Lys Thr Pro Lys Asn Gln Glu Ala Leu Arg Met Ala Asn Val Ala Glu 305 310 315 Asn Ser Ser Ser Asp Gln Arg Gln Ala Cys Lys His Glu Leu Tyr Val Ser Phe Arg Asp Leu Gly Trp Gln Asp Trp Ile Ile Ala Pro Glu 340 345 Gly Tyr Ala Ala Tyr Tyr Cys Glu Gly Glu Cys Ala Phe Pro Leu Asn 355 Ser Tyr Met Asn Ala Thr Asn His Ala Ile Val Gln Thr Leu Val His Phe Ile Asn Pro Glu Thr Val Pro Lys Pro Cys Cys Ala Pro Thr Gln 385 390 395 Leu Asn Ala Ile Ser Val Leu Tyr Phe Asp Asp Ser Ser Asn Val Ile Leu Lys Lys Tyr Arg Asn Met Val Val Arg Ala Cys Gly Cys His

- (2) INFORMATION FOR SEQ ID NO:3:
 - (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 102 amino acids
 (B) TYPE: amino acid
 (C) STRANDEDNESS:
 (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: protein
 - (ix) FEATURE:
 - (A) NAME/KEY: Protein
 - (B) LOCATION: 1..102

(D) OTHER INFORMATION: /label= OPX /note= "wherein each Xaa is independently selected from a group of one or more specified amino acids as defined in the specification."

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:3:

Cys Xaa Xaa His Glu Leu Tyr Val Xaa Phe Xaa Asp Leu Gly Trp Xaa 1 10 15

Asp Trp Xaa Ile Ala Pro Xaa Gly Tyr Xaa Ala Tyr Tyr Cys Glu Gly 25

Glu Cys Xaa Phe Pro Leu Xaa Ser Xaa Met Asn Ala Thr Asn His Ala 35

Ile Xaa Gln Xaa Leu Val His Xaa Xaa Xaa Pro Xaa Xaa Val Pro Lys

Xaa Cys Cys Ala Pro Thr Xaa Leu Xaa Ala Xaa Ser Val Leu Tyr Xaa

5

65

75

80

Asp Xaa Ser Xaa Asn Val Xaa Leu Xaa Lys Xaa Arg Asn Met Val Val 85 90 95

Xaa Ala Cys Gly Cys His 100

(2) INFORMATION FOR SEQ ID NO:4:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 97 amino acids
 (B) TYPE: amino acid
 (C) STRANDEDNESS:

 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein

(1x) FEATURE:

(A) NAME/KEY: Protein

(B) LOCATION: 1..97

(D) OTHER INFORMATION: /label= Generic-Seq-7

/note= "wherein each Xaa is independently selected from a group of one or more specified amino acids as defined in the specification."

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:4:

Leu Xaa Xaa Xaa Phe Xaa Xaa Xaa Gly Trp Xaa Xaa Xaa Xaa Xaa Xaa 1

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Asn His Ala Xaa Xaa Xaa Xaa Xaa 40

Val Xaa Leu Xaa Xaa Xaa Xaa Xaa Met Xaa Val Xaa Xaa Cys Xaa Cys 90 95

Xaa

(2) INFORMATION FOR SEQ ID NO:5:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 102 amino acids
 (B) TYPE: amino acid
 (C) STRANDEDNESS:

 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
- (1X) FEATURE:

 (A) NAME/KEY: Protein

 (B) LOCATION: 1..102

 (D) OTHER INFORMATION: /label= Generic-Seq-8

 /note= "wherein each Xaa is independently selected from a group of one or more specified amino acids as defined in the specification."
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:
 - Cys Xaa Xaa Xaa Leu Xaa Xaa Xaa Phe Xaa Xaa Gly Trp Xaa 1 5 10
 - Xaa Xaa Xaa Xaa Pro Xaa Xaa Xaa Ala Xaa Tyr Cys Xaa Gly
 - Xaa Cys Xaa Xaa Pro Xaa Xaa Xaa Xaa Xaa Xaa Xaa Asn His Ala 35 40

Xaa Cys Cys Xaa Pro Xaa Xaa Xaa Xaa Xaa Xaa Xaa Leu Xaa Xaa 65 70 75

Xaa Xaa Xaa Xaa Xaa Val Xaa Leu Xaa Xaa Xaa Xaa Met Xaa Val

Xaa Xaa Cys Xaa Cys Xaa 100

- (2) INFORMATION FOR SEQ ID NO:6:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 97 amino acids (B) TYPE: amino acid

 - STRANDEDNESS: single
 - TOPOLOGY: linear
 - (ii) MOLECULE TYPE: protein
 - (ix) FEATURE:
 - (A) NAME/KEY: Protein (B) LOCATION: 1..97
 - (D) OTHER INFORMATION: /label= Generic-Seq-9

/note= "wherein each Xaa is independently selected from a group of one or more specified amino acids as defined in the specification."

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:

Pro Xaa Xaa Xaa Xaa Xaa Xaa Cys Xaa Gly Xaa Cys Xaa Xaa Xaa Xaa 20 30

Xaa

- (2) INFORMATION FOR SEQ ID NO:7:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 102 amino acids (B) TYPE: amino acid

 - STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: protein
 - (ix) FEATURE:
 - (A) NAME/KEY: Protein (B) LOCATION: 1..102
 - (D) OTHER INFORMATION: /label= Generic-Seq-10 /note= "wherein each Xaa is independently selected from a group of one or more specified amino acids as defined in the specification."
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:7:

Xaa Xaa Xaa Xaa Aaa Aaa Xaa Aaa Xaa Cys Xaa Gly 20 25 30

Xaa Xaa Cys Xaa Pro Xaa Xaa Xaa Xaa Xaa Xaa Xaa Leu Xaa Xaa 65 70 80

Xaa Xaa Cys Xaa Cys Xaa 100

- (2) INFORMATION FOR SEQ ID NO:8:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 5 amino acids
 (B) TYPE: amino acid

 - (C) STRANDEDNESS: (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:

- (1X) FEATURE:

 (A) NAME/KEY: Protein

 (B) LOCATION: 1..5

 (D) OTHER INFORMATION: /note= "wherein each Xaa is independently selected from a group of one or more specified amino acids as defined in the specification"
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:8:

Cys Xaa Xaa Xaa Xaa 1

- (2) INFORMATION FOR SEQ ID NO:9:
 - (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 5 amino acids
 (B) TYPE: amino acid
 (C) STRANDEDNESS:

 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (1X) FEATURE:

 (A) NAME/KEY: Protein

 (B) LOCATION: 1..5

 (D) OTHER INFORMATION: /note= "wherein each Xaa is independently selected from a group of one or more specified amino acids as defined in the specification"
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:9:

Cys Xaa Xaa Xaa Xaa 1